

Claim Amendments

Please cancel claims 21, 23, and 27 without prejudice, and please amend claims 1, 4-10, 13, 15, 18, 21, 22, 26, 32-34, 36, 37, and 40-42, as follows. This listing of claims will replace all prior versions and listings of claims in the instant application.

Listing of Claims:

1. (Currently Amended) An antimicrobial sulfonamide derivative, or a salt or a hydrate thereof, comprising:

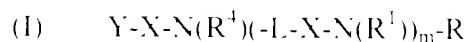
a core cyclic peptide or core antibiotic of a lipopeptide antibiotic; and
a lipophilic moiety,

wherein said lipophilic moiety is covalently attached to the core cyclic peptide or core ~~eyelie~~-antibiotic *via* a linking chain which includes a sulfonamide linkage and wherein said core cyclic peptide or core antibiotic is not of laspartomycin or polymyxin.

2. (Original) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 in which the linking chain is a sulfonamide linkage.

3. (Original) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 in which the linking chain is a linker that links the core cyclic peptide or core antibiotic to the lipophilic moiety.

4. (Currently Amended) The antimicrobial sulfonamide derivative, salt or hydrate of Claim 1 which is a compound according to structural Formula (I):



wherein:

Y is a lipophilic moiety;

Each each X is independently selected from the group consisting of -co-SO₂-, -CS-, -PO-, -OP(O)-, -OC(O)-, -NHCO and N(R¹)CO-N(R¹)CO- with the proviso that at least one X is -SO₂-;

M is 0 or 1;

L is a linker;

N is nitrogen;

R¹ and R⁴ are each independently selected from the group consisting of hydrogen, (C₁-C₂₅) alkyl optionally substituted with one or more of the same or different R² groups, (C₁-C₂₅) heteroalkyl optionally substituted with one or more of the same or different R² groups, (C₅-C₃₀) aryl optionally substituted with one or more of the same or different R² groups, (C₅-C₃₀) arylaryl optionally substituted with one or more of the same or different R² groups, (C₅-C₃₀) biaryl optionally substituted with one or more of the same or different R² groups, five to thirty membered heteroaryl optionally substituted with one or more of the same or different R² groups, (C₆-C₃₀) arylalkyl optionally substituted with one or more of the same or different R₂ groups and six to thirty membered heteroarylalkyl optionally substituted with one or more of the same or different R₂ groups;

each R² is independently selected from the group consisting of -OR³, -SR³, -NR³R³, -CN, -NO₂, -N³, -C(O)OR³, -C(O)NR³R³, -C(S)NR³R³, -C(NR³)NR³R³, -CHO, -R³CO, -SO₂R³, -SOR³, -PO(OR³)₂, -PO(OR³), -CO₂H, -SO₃H, -PO₃H, halogen and trihalomethyl;

each R³ is independently selected from the group consisting of hydrogen, (C₁-C₆) alkyl, (C₅-C₁₀) aryl, five to sixteen membered heteroaryl, (C₆-C₁₆) arylalkyl and six to sixteen membered heteroarylalkyl; and

R is a core cyclic peptide or core antibiotic of a lipopeptide antibiotic, wherein said core cyclic peptide or core antibiotic is not of laspartomycin or polymyxin.

5. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of laspartomycin, zaomycin, crystallomycin, aspartocin,

amphomycin, glumamycin, brevistin, cerexin A, cerexin B, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145, Antibiotic A-21978C or tsushimaycin.

6. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of ~~laspartomyein~~—zaomycin, crystallomycin, aspartocin, amphomycin, glumamycin, brevistin, cerexin A, cerexin B, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145, Antibiotic A-21978C or tsushimaycin.

7. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of ~~laspartomyein~~—aspartocin, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A-54145 or Antibiotic A-21978C.

8. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of ~~laspartomyein~~, aspartocin, Antibiotic A-30912, Antibiotic A-1437, Antibiotic A54145 or Antibiotic A-21978C.

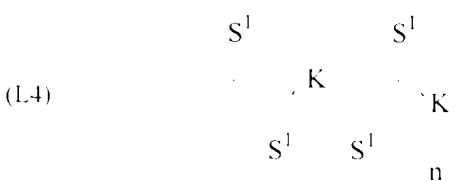
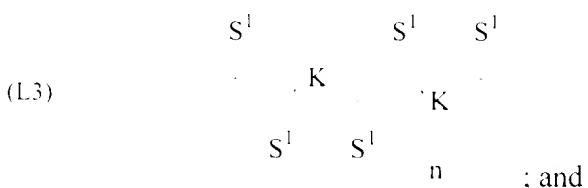
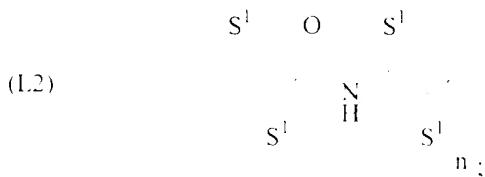
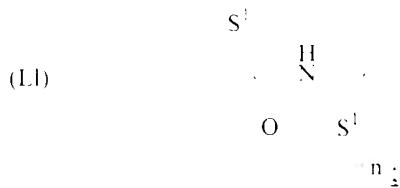
9. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core cyclic peptide of ~~laspartomyein~~ or aspartocin.

10. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 4 in which R is the core antibiotic of ~~laspartomyein~~ or aspartocin.

11. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which m is 1.

12. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which R¹ and R⁴ are hydrogen.

13. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 4 in which L is selected from the group consisting of:



or a pharmaceutically acceptable salt or hydrate thereof, wherein:

n is 0, 1, 2 or 3;

each S¹ is independently selected from the group consisting of hydrogen, (C₁ – C₁₀) alkyl optionally substituted with one or more of the same or different R⁵ groups, (C₁ – C₁₀) heteroalkyl optionally substituted with one or more of the same or different R⁵ groups, (C₅ – C₁₀) aryl optionally substituted with one or more of the same or different R⁵ groups, (C₅–C₁₅) arylaryl optionally substituted with one or more of the same or different R⁵ groups, (C₅–C₁₅) biaryl optionally substituted with one or more of the same or different R⁵ groups, five to ten membered heteroaryl optionally substituted with one or more of the same or different R⁵ groups, (C₆–C₁₆) arylalkyl optionally substituted with one or more of the same or different R⁵ groups and six to sixteen membered heteroarylalkyl optionally substituted with one or more of the same or different R⁵ groups;

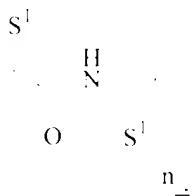
each R⁵ is independently selected from the group consisting of -OR⁶, -SR⁶, -NR⁶R⁶, -CN, -NO₂, -N₃, -C(O)OR⁶, -C(O)NR⁶R⁶, -C(S)NR⁶R⁶, -C(NR⁶)NR⁶R⁶, -CHO, -R⁶CO, -SO₂R⁶, -SOR⁶, -PO(OR⁶)₂, -PO(OR⁶), -CO₂H, -SO₃H, -PO₃H, halogen and trihalomethyl;

each R⁶ is independently selected from the group consisting of hydrogen, (C₁-C₆) alkyl, (C₅-C₁₀) aryl, five to sixteen membered heteroaryl, (C₆-C₁₆) arylalkyl and six to sixteen membered heteroarylalkyl; and

each K is independently selected from the group consisting of oxygen, nitrogen and sulfur.

14. (Original) The antimicrobial sulfonamide of Claim 13 in which each S¹ is independently a side-chain of a genetically encoded α -amino acid.

15. (Currently Amended) The antimicrobial sulfonamide of Claim 13 in which L is:



16. (Original) The antimicrobial sulfonamide derivative of Claim 15 in which each S¹ is independently a side-chain of a genetically encoded α -amino acid.

17. (Original) The antimicrobial sulfonamide derivative of Claim 15 in which n is 0.

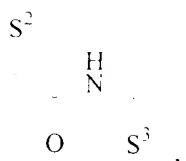
18. (Currently Amended) The ~~compound~~—antimicrobial sulfonamide derivative of Claim 17 in which S¹ is hydrogen, X²-Y is decan-yl and R is the core cyclic peptide of aspartocin.

19. (Original) The antimicrobial sulfonamide derivative of Claim 17 in which S¹ is -CH₂-CO₂H, -CH₂-CH₂-CO₂H, -C(OH)H-CONH₂, -CH₂-CONH₂ or -CH₂-CH₂-CONH₂ or a salt or hydrate thereof.

20. (Original) The antimicrobial sulfonamide derivative of Claim 17 in which S¹ is -CH₂-indol-2-yl or -CH₂-phenyl.

21. (Cancelled)

22. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 13 in which L is:



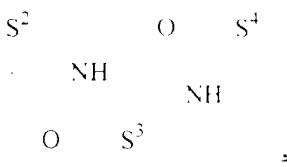
wherein S² and S³ are each independently a side chain of a genetically encoded α -amino acid.

23. (Cancelled)

24. (Original) The antimicrobial sulfonamide derivative of Claim 22 in which S² is hydrogen, -CH₂-indol-2-yl, -CH₂-CONH₂ or -CH₂-CH₂-CONH₂ and S³ is -CH₂-CO₂H, -CH₂-CH₂-CO₂H or a salt or hydrate thereof.

25. (Original) The antimicrobial sulfonamide derivative of Claim 22 in which S² is -CH₂-CO₂H, -CH₂-CH₂-CO₂H or a salt or hydrate thereof and S³ is -C(OH)H-CONH₂.

26. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 13 in which L is:



wherein S^2 , S^3 , and S^4 are each independently a side chain of a genetically encoded α -amino acid.

27. (Cancelled)

28. (Original) The antimicrobial sulfonamide derivative of Claim 26 in which S^2 is -CH₂-indol-2-yl, S^3 is -CH₂-CONH₂ or -CH₂-CH₂-CONH₂ and S^4 is -CH₂-CO₂H, -CH₂-CH₂-CO₂H or a salt or hydrate thereof.

29. (Original) The antimicrobial sulfonamide derivative of Claim 26 in which S^2 is -CH₂-indol-2-yl, S^3 is -CH₂-CO₂H, CH₂-CH₂-CO₂H or a salt or hydrate thereof and S^4 is -CH₂-CONH₂, -CH₂-CH₂-CONH₂ or -C(OH)H-CONH₂.

30. (Original) The antimicrobial sulfonamide derivative of Claim 4 in which m is 0.

31. (Original) The antimicrobial sulfonamide derivative of Claim 30 in which R⁴ is hydrogen.

32. (Currently Amended) The antimicrobial sulfonamide derivative of Claim 30 in which R is the core antibiotic of ~~laspantomyein~~ or aspartocin.

33. (Currently Amended) The antimicrobial sulfonamide derivative of ~~Claim 32~~ Claim 30 in which R is the core cyclic peptide of ~~laspantomyein~~ or aspartocin.

34. (Currently Amended) A pharmaceutical composition comprising a compound an antimicrobial sulfonamide derivative according to Claim 4 and a pharmaceutically acceptable adjuvant, excipient, carrier or diluent.

35. (Original) A method for treating or preventing a microbial infection, said method comprising the step of administering to a subject a therapeutically effective amount of a compound according to Claim 4 or a therapeutically effective amount of a pharmaceutical composition according to Claim 34.

36. (Currently Amended) A method of inhibiting microbial growth, said method comprising the step of administering to a microbe an antimicrobially effective amount of a compound an antimicrobial sulfonamide derivative according to Claim 4 or an antimicrobially effective amount of a pharmaceutical composition according to Claim 34.

37. (Currently Amended) A method for making an antimicrobial sulfonamide derivative comprising sulfonylating an-a core antibiotic or core cyclic peptide with a lipophilic sulfonyl derivative, thereby providing a-an antimicrobial sulfonamide derivative.

38. (Original) The method of Claim 37 in which the lipophilic sulfonyl derivative is a activated lipophilic sulfonyl ester or a lipophilic sulfonyl halide.

39. (Original) The method of Claim 38 in which the activated lipophilic sulfonyl ester is a lipophilic hydroxybenzotriazole ester.

40. (Currently Amended) The method of ~~Claim 39~~Claim 38 in which the lipophilic sulfonyl halide is a lipophilic sulfonyl chloride.

41. (Currently Amended) A method for making an antimicrobial sulfonamide derivative comprising:

sulfonylating a linker with a lipophilic sulfonyl compound, thereby providing a lipophilic sulfonamide linker; and

covalently attaching the lipophilic sulfonamide linker to ~~an-a~~ core antibiotic or core cyclic peptide wherein said core cyclic peptide or core antibiotic is not of polymyxin, thereby yielding ~~a-an~~ antimicrobial sulfonamide derivative.

42. (Currently Amended) A method for making an antimicrobial sulfonamide derivative comprising:

covalently attaching a linker to ~~an-a~~ core antibiotic or core cyclic peptide, thereby providing an linker core antibiotic or linker core cyclic peptide; and

sulfonylating the linker core antibiotic or linker core cyclic peptide with a lipophilic sulfonyl derivative, thereby yielding ~~a-an~~ antimicrobial sulfonamide derivative.